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(FILE 'USPAT' ENTERED AT 14:01:29 ON 30 SEP 92)

L1 9 S CAPSULAR POLYMER#
L2 1112 S REDUCTIVE AMINATION
L3 11560 S IMMUNOGEN? OR ANTIGEN?
L4 1182 S BACTERIAL AND (TOXIN# OR TOXOID#)
L5 19 S L4 AND L3 AND L2
L6 6 S L5 AND L1

=>

APS
07/819305

File APS

07/819305

s capsular polymer#
909 CAPSULAR
168344 POLYMER#
L1 9 CAPSULAR POLYMER#
(CAPSULAR(W)POLYMER#)

=> d 1-9

✓ 1. 5,097,020, Mar. 17, 1992, Immunogenic conjugates; Porter W. Anderson, et al., 530/403; 424/88, 89, 92; 435/68.1; 530/395, 402, 404, 405, 406, 807 [IMAGE AVAILABLE]

2. 5,077,320, Dec. 31, 1991, Microvoid-containing polymer particles; Hideki Touda, et al., 521/65 [IMAGE AVAILABLE]

3. 4,972,000, Nov. 20, 1990, Hollow polymer particles, process for production thereof, and use thereof as pigment; Nobuo Kawashima, et al., 521/54; 427/222; 428/407; 521/55, 57, 134; 525/902 [IMAGE AVAILABLE]

✓ 4. 4,902,506, Feb. 20, 1990, Immunogenic conjugates; Porter W. Anderson, et al., 424/92, 88; 530/350

5. 4,883,757, Nov. 28, 1989, Bioemulsifier production by Acinetobacter calcoaceticus strains; David L. Gutnick, et al., 435/252.1; 252/351; 435/253.6, 822 [IMAGE AVAILABLE]

6. 4,808,700, Feb. 28, 1989, Immunogenic conjugates of non-toxic E. coli LT-B enterotoxin subunit and **capsular** **polymers**;; Porter W. Anderson, et al., 424/92; 435/6, 172.3, 240.27; 514/12; 530/403, 807, 808, 812; 935/12

✓ 7. 4,762,713, Aug. 9, 1988, Boosting of immunogenic conjugate vaccinations by unconjugated bacterial **capsular** **polymers**;; Porter W. Anderson, 424/92, 88

✓ 8. 4,761,283, Aug. 2, 1988, Immunogenic conjugates; Porter W. Anderson, 424/92, 88; 530/350

✓ 9. 4,673,574, Jun. 16, 1987, Immunogenic conjugates; Porter W. Anderson, 424/92; 530/350

=> s ~~polymer~~ amination
6226 REDUCTIVE
2943 AMINATION
L2 1112 REDUCTIVE AMINATION
(REDUCTIVE(W)AMINATION)

=> s immunogen? or antigen?
2626 IMMUNOGEN?
11137 ANTIGEN?
L3 11560 IMMUNOGEN? OR ANTIGEN?

=> s bacterial and (toxin# or toxoid#)
20699 BACTERIAL
2761 TOXIN#
474 TOXOID#
L4 1182 BACTERIAL AND (TOXIN# OR TOXOID#)

=> s 14 and 13 and 12
L5 19 L4 AND L3 AND L2

=> s 15 and 11
L6 6 L5 AND L1

=> d 1=0

1. 5,097,020, Mar. 17, 1992, **Immunogenic** conjugates; Porter W. Anderson, et al., 530/403; 424/88, 89, 92; 435/68.1; 530/395, 402, 404, 405, 406, 807 [IMAGE AVAILABLE]
2. 4,902,506, Feb. 20, 1990, **Immunogenic** conjugates; Porter W. Anderson, et al., 424/92, 88; 530/350
3. 4,808,700, Feb. 28, 1989, **Immunogenic** conjugates of non-toxic E. coli LT-B enterotoxin subunit and **capsular** **polymers**;; Porter W. Anderson, et al., 424/92; 435/6, 172.3, 240.27; 514/12; 530/403, 807, 808, 812; 935/12
4. 4,762,713, Aug. 9, 1988, Boosting of **immunogenic** conjugate vaccinations by unconjugated **bacterial** **capsular** **polymers**;; Porter W. Anderson, 424/92, 88
5. 4,761,283, Aug. 2, 1988, **Immunogenic** conjugates; Porter W. Anderson, 424/92, 88; 530/350
6. 4,673,574, Jun. 16, 1987, **Immunogenic** conjugates; Porter W. Anderson, 424/92; 530/350

=> d bib

US PAT NO: 5,097,020 [IMAGE AVAILABLE] L6: 1 of 6
 DATE ISSUED: Mar. 17, 1992
 TITLE: **Immunogenic** conjugates
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 Ronald J. Eby, Rochester, NY
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 APPL-NO: 07/423,081
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=> d &lms

US PAT NO: 5,097,020 [IMAGE AVAILABLE] L6: 1 of 6

CLAIMS:

CLMS(1)

We claim:

1. An **immunogenic** conjugate, comprising: the **reductive**
 amination product of a **capsular** **polymer** fragment having at
 least two carbonyl groups and derived from the **capsular** **polymer**
 of a **bacterial** pathogen by a process which comprises first treating
 said polymer with acid, base or enzyme and then generating carbonyl
 groups by treatment with an oxidizing agent, and a **bacterial**
 toxin or **toxoid**, said conjugate comprising a cross-linked
 conjugate in which there is a direct covalent linkage between the
 capsular **polymer** fragment and the **toxin** or **toxoid**.

CLMS(2)

2. The **immunogenic** conjugate of claim 1, wherein the **capsular**
 polymer is **immunogenic** in mature humans and less **immunogenic**
 in infant humans.

CLMS(3)

3. The ****immunogenic**** conjugate of claim 1, wherein the ****reductive**** ****amination**** is performed in the presence of cyanoborohydride anions.

CLMS(4)

4. The ****immunogenic**** conjugate of claim 1, wherein the ****toxin**** or ****toxoid**** is diphtheria ****toxin**** or ****toxoid****.

CLMS(5)

5. The ****immunogenic**** conjugate of claim 4, wherein the ****toxoid**** is CRM.sub.197.

CLMS(6)

6. The ****immunogenic**** conjugate of claim 1, wherein the ****toxin**** or ****toxoid**** is tetanus ****toxin**** or ****toxoid****.

CLMS(7)

7. The ****immunogenic**** conjugate of claim 1, wherein the ****toxin**** or ****toxoid**** is a pseudomonas ****toxin**** or ****toxoid****.

CLMS(8)

8. The ****immunogenic**** conjugate of claim 1, wherein the ****toxin**** or ****toxoid**** is a staphylococcus ****toxin**** or ****toxoid****.

CLMS(9)

9. The ****immunogenic**** conjugate of claim 1, wherein the ****toxin**** or ****toxoid**** is a streptococcus ****toxin**** or ****toxoid****.

CLMS(10)

10. The ****immunogenic**** conjugate of claim 1, wherein the ****toxin**** or ****toxoid**** is pertussis ****toxin**** or ****toxoid****.

CLMS(11)

11. The ****immunogenic**** conjugate of claim 1, wherein the ****toxin**** or ****toxoid**** is an Escherichia coli ****toxin**** or ****toxoid****.

CLMS(12)

12. The ****immunogenic**** conjugate of claim 1, wherein the ****bacterial**** pathogen is Haemophilus influenzae type b.

CLMS(13)

13. The ****immunogenic**** conjugate of claim 1, wherein the ****bacterial**** pathogen is Escherichia coli.

CLMS(14)

14. The ****immunogenic**** conjugate of claim 1, wherein the ****bacterial**** pathogen is Neisseria meningitidis.

CLMS(15)

15. The ****immunogenic**** conjugate of claim 1, wherein the ****bacterial**** pathogen is Neisseria meningitidis serogroup A.

CLMS(16)

16. The ****immunogenic**** conjugate of claim 1, wherein the ****bacterial**** pathogen is *Neisseria meningitidis* serogroup C.

CLMS(17)

17. The ****immunogenic**** conjugate of claim 1, wherein the ****bacterial**** pathogen is *Streptococcus pneumoniae*.

CLMS(18)

18. The ****immunogenic**** conjugate of claim 1, wherein the ****bacterial**** pathogen is *Streptococcus pneumoniae* serotype 6.

CLMS(19)

19. The ****immunogenic**** conjugate of claim 1, wherein the ****bacterial**** pathogen is *Streptococcus pneumoniae* serotype 12.

CLMS(20)

20. The ****immunogenic**** conjugate of claim 1, wherein the ****bacterial**** pathogen is *Streptococcus pneumoniae* serotype 14.

CLMS(21)

21. The ****immunogenic**** conjugate of claim 1, wherein the ****bacterial**** pathogen is *Streptococcus pneumoniae* serotype 19.

CLMS(22)

22. The ****immunogenic**** conjugate of claim 1, wherein the ****bacterial**** pathogen is *Streptococcus pneumoniae* serotype 23.

CLMS(23)

23. The ****immunogenic**** conjugate of claim 1, wherein the ****bacterial**** pathogen is *Streptococcus pneumoniae* serotype 51.

CLMS(24)

24. The ****immunogenic**** conjugate of claim 5, wherein the ****bacterial**** pathogen is *Haemophilis influenzae* type b.

CLMS(25)

25. The ****immunogenic**** conjugate of claim 5, wherein the ****bacterial**** pathogen is *Streptococcus pneumoniae* serotype 6.

CLMS(26)

26. The ****immunogenic**** conjugate of claim 5, wherein the ****bacterial**** pathogen is *Streptococcus pneumoniae* serotype 14.

CLMS(27)

27. The ****immunogenic**** conjugate of claim 5, wherein the ****bacterial**** pathogen is *Streptococcus pneumoniae* serotype 19.

CLMS(28)

28. The ****immunogenic**** conjugate of claim 5, wherein the ****bacterial**** pathogen is *Streptococcus pneumoniae* serotype 23.

CLMS(29)

29. The ****immunogenic**** conjugate of claim 1, wherein the fragment is

derived from the **capsular** **polymer** by oxidative cleavage.

CLMS(30)

30. The **immunogenic** conjugate of claim 1, wherein the fragment is derived from the **capsular** **polymer** by periodate.

CLMS(31)

31. An **immunogenic** conjugate comprising: a formalin treated **reductive** **amination** product of a **capsular** **polymer** fragment having at least two (2) carbonyl groups and derived from the **capsular** **polymer** of a **bacterial** pathogen by a process which comprises first treating said polymer with acid, base or enzyme and then generating carbonyl groups by treatment with an oxidizing agent, and a **bacterial** **toxin** or **toxoid**, said conjugate comprising a cross-linked conjugate in which there is a direct covalent linkage between the **capsular** **polymer** fragment and the **toxin** or **toxoid**.

CLMS(32)

32. The **immunogenic** conjugate of claim 31 wherein the **bacterial** **toxoid** is diphtheria **toxoid**.

CLMS(33)

33. The **immunogenic** conjugate of claim 31 wherein the **toxoid** is CRM.sub.197.

CLMS(34)

34. The **immunogenic** conjugate of claim 31 wherein the **bacterial** **toxin** or **toxoid** is tetanus **toxin** or **toxoid**.

CLMS(35)

35. A method for preparing an **immunogenic** conjugate, comprising forming the **reductive** **amination** product of a **capsular** **polymer** fragment having at least two carbonyl groups and derived from the **capsular** **polymer** of a **bacterial** pathogen by a process which comprises first treating said polymer with acid, base or enzyme and then generating carbonyl groups by treatment with an oxidizing agent, and a **bacterial** **toxin** or **toxoid**, said **reductive** **amination** being performed in the presence of cyanborohydride ions, said conjugate comprising a cross-linked conjugate in which there is a direct covalent linkage between the **capsular** **polymer** fragment and the **toxin** or **toxoid**.

CLMS(36)

36. The method of claim 35, wherein the **capsular** **polymer** is **immunogenic** in mature humans and less **immunogenic** in infant humans.

CLMS(37)

37. The method of claim 35, wherein the **toxin** or **toxoid** is diphtheria **toxin** or **toxoid**.

CLMS(38)

38. The method of claim 35, wherein the **toxin** or **toxoid** is CRM.sub.197.

CLMS(39)

39. The method of claim 35, wherein the **toxin** or **toxoid** is tetanus **toxin** or **toxoid**.

CLMS(40)

40. The method of claim 35, wherein the **toxin** or **toxoid** is pseudomonas **toxin** or **toxoid**.

CLMS(41)

41. The method of claim 35, wherein the **toxin** or **toxoid** is staphylococcus **toxin** or **toxoid**.

CLMS(42)

42. The method of claim 35, wherein the **toxin** or **toxoid** is streptococcus **toxin** or **toxoid**.

CLMS(43)

43. The method of claim 35, wherein the **toxin** or **toxoid** is pertussis **toxin** or **toxoid**.

CLMS(44)

44. The method of claim 35, wherein the **toxin** or **toxoid** is an Escherichia coli **toxin** or **toxoid**.

CLMS(45)

45. The method of claim 35, wherein the pathogen is Haemophilus influenzae type b.

CLMS(46)

46. The method of claim 35, wherein the pathogen is Escherichia coli.

CLMS(47)

47. The method of claim 35, wherein the pathogen is Neisseria meningitidis.

CLMS(48)

48. The method of claim 35, wherein the pathogen is Streptococcus pneumoniae.

CLMS(49)

49. The method of claim 35, wherein the pathogen is Pseudomonas.

CLMS(50)

50. The method of claim 37, wherein the pathogen is Haemophilus influenzae b.

CLMS(51)

51. The method of claim 37, wherein the pathogen is Streptococcus pneumonia.

CLMS(52)

52. The method of claim 35, wherein the fragment is derived from the .
capsular **polymer** by oxidative cleavage.

CLMS(53)

53. A method for preparing an **immunogenic** conjugate, comprising: forming the **reductive** **amination** product of a **capsular** **polymer** fragment having at least two carbonyl groups and derived from the **capsular** **polymer** of a **bacterial** pathogen, and a **bacterial** **toxin** or **toxoid**, said **reductive** **amination** being performed in the presence of cyanoborohydride ions, wherein the fragment is derived from the **capsular** polymer by periodate said conjugate comprising a cross-linked conjugate.

CLMS(54)

54. A method for preparing an immunogenic conjugate comprising: forming the reductive amination product of a capsular polymer fragment having at least two carbonyl groups and derived from the capsular polymer of a bacterial pathogen, and a bacterial toxin or toxoid, said reductive amination being performed in the presence of cyanohorohydride ions, in which the fragment is produced from the capsular polymer by first treating said polymer with acid, base or enzyme and then generating carbonyl groups by treatment with an oxidizing agent, said conjugate comprising a crosslinked conjugate.

CLMS(55)

55. The method of claim 54, further comprising treating said reductive amination product with formalin.

CLMS(56)

56. The method of claim 55, wherein the bacterial toxoid is diphtheria toxoid.

CLMS(57)

57. The method of claim 55, wherein the toxoid is CRM.sub.197.

CLMS(58)

58. The method of claim 55, wherein the bacterial toxin or toxoid is tetanus toxin or toxoid.

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US PAT NO: 4,902,506

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CLAIMS:

CLMS(1)

We claim:

1. An **immunogenic** conjugate comprising the **reductive** **amination** product of a **capsular** **polymer** fragment having a chain length of from about 10 to about 30 monomeric units and at least two carbonyl groups, which fragment is derived from the **capsular** **polymer** of a Streptococcus pneumoniae or Haemophilus influenzae bacterium, and a **bacterial** **toxin** or **toxoid**, said conjugate comprising a crosslinked conjugate.

CLMS(2)

2. The **immunogenic** conjugate of claim 1, wherein the **capsular** **polymer** is **immunogenic** in mature humans and less **immunogenic** in infant humans.

CLMS(3)

3. The **immunogenic** conjugate of claim 1, wherein the **reductive amination** is performed in the presence of cyanoborohydride anions.

CLMS(4)

4. The **immunogenic** conjugate of claim 1, wherein the **toxin** or **toxoid** is diphtheria **toxin** or **toxoid**.

CLMS(5)

5. The **immunogenic** conjugate of claim 1, wherein the **toxoid** is CRM.sub.197.

CLMS(6)

6. The **immunogenic** conjugate of claim 1, wherein the **toxin** or **toxoid** is tetanus **toxin** or **toxoid**.

CLMS(7)

7. The **immunogenic** conjugate of claim 1, wherein the **toxin** or **toxoid** is a pseudomonas **toxin** or **toxoid**.

CLMS(8)

8. The **immunogenic** conjugate of claim 1, wherein the **toxin** or **toxoid** is a staphylococcus **toxin** or **toxoid**.

CLMS(9)

9. The **immunogenic** conjugate of claim 1, wherein the **toxin** or **toxoid** is a streptococcus **toxin** or **toxoid**.

CLMS(10)

10. The **immunogenic** conjugate of claim 1, wherein the **toxin** or **toxoid** is pertussis **toxin** or **toxoid**.

CLMS(11)

11. The **immunogenic** conjugate of claim 1, wherein the **toxin** or **toxoid** is an Escherichia coli **toxin** or **toxoid**.

CLMS(12)

12. The **immunogenic** conjugate of claim 1, wherein the **bacterial** pathogen is Haemophilus influenzae type b.

CLMS(13)

13. The **immunogenic** conjugate of claim 1, wherein the **bacterial** pathogen is Streptococcus pneumoniae serotype 3.

CLMS(14)

14. The **immunogenic** conjugate of claim 1, wherein the **bacterial** pathogen is Streptococcus pneumoniae serotype 6.

CLMS(15)

15. The **immunogenic** conjugate of claim 1, wherein the **bacterial** pathogen is Streptococcus pneumoniae serotype 12.

CLMS(16)

16. The ****immunogenic**** conjugate of claim 1, wherein the ****bacterial**** pathogen is *Streptococcus pneumoniae* serotype 14.

CLMS(17)

17. The ****immunogenic**** conjugate of claim 1, wherein the ****bacterial**** pathogen is *Streptococcus pneumoniae* serotype 19.

CLMS(18)

18. The ****immunogenic**** conjugate of claim 1, wherein the ****bacterial**** pathogen is *Streptococcus pneumoniae* serotype 23.

CLMS(19)

19. The ****immunogenic**** conjugate of claim 1, wherein the ****bacterial**** pathogen is *Streptococcus pneumoniae* serotype 51.

CLMS(20)

20. The ****immunogenic**** conjugate of claim 5, wherein the ****bacterial**** pathogen is *Haemophilis influenzae* type b.

CLMS(21)

21. The ****immunogenic**** conjugate of claim 5, wherein the ****bacterial**** pathogen is *Streptococcus pneumoniae* serotype 6.

CLMS(22)

22. The ****immunogenic**** conjugate of claim 5, wherein the ****bacterial**** pathogen is *Streptococcus pneumoniae* serotype 14.

CLMS(23)

23. The ****immunogenic**** conjugate of claim 5, wherein the ****bacterial**** pathogen is *Streptococcus pneumoniae* serotype 19.

CLMS(24)

24. The ****immunogenic**** conjugate of claim 5, wherein the ****bacterial**** pathogen is *Streptococcus pneumoniae* serotype 23.

CLMS(25)

25. The ****immunogenic**** conjugate of claim 1, wherein the fragment is derived from the ****capsular**** ****polymer**** by oxidative cleavage.

CLMS(26)

26. The ****immunogenic**** conjugate of claim 1, wherein the fragment is derived from the ****capsular**** ****polymer**** by periodate.

CLMS(27)

27. An ****immunogenic**** conjugate comprising: a formalin treated ****reductive**** ****amination**** product of a ****capsular**** ****polymer**** fragment having a chain length of from about 10 to about 30 monomeric units and at least two carbonyl groups, which fragment is derived from the ****capsular**** ****polymer**** of a *Streptococcus pneumoniae* or *Haemophilus influenzae* bacterium, and a ****bacterial**** ****toxin**** or ****toxoid****, said conjugate comprising a crosslinked conjugate.

CLMS(28)

28. The ****immunogenic**** conjugate of claim 27, wherein the ****bacterial**** ****toxoid**** is diphtheria ****toxoid****.

CLMS(29)

29. The ****immunogenic**** conjugate of claim 27, wherein the ****toxoid**** is CRM.sub.197.

30. The ****immunogenic**** conjugate of claim 27, wherein the ****bacterial**** ****toxin**** or ****toxoid**** is tetanus ****toxin**** or ****toxoid****.

CLMS(31)

31. The ****immunogenic**** conjugate of claim 1, in which the fragment is produced from ****capsular**** ****polymer**** of Streptococcus pneumoniae by first treating said polymer with acid, base or enzyme and then generating aldehyde groups by treatment with an oxidizing agent.

CLMS(32)

32. The ****immunogenic**** conjugate of claim 1, in which the fragment is produced from ****capsular**** ****polymer**** of Haemophilus influenzae by first treating said polymer with acid, base or enzyme and then generating aldehyde groups by treatment with an oxidizing agent.

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CLAIMS:

CLMS(1)

We claim:

1. A conjugate of a PRP polysaccharide fragment, having reducing terminal groups derived from the capsular polysaccharide of Haemophilus influenzae type b by selective acidic hydrolysis of a portion of the ribosyl ribitol linkages therein, and a ****bacterial**** binding subunit, which binding subunit is a non-toxic polypeptide, having one or more immunoreactive and ****antigenic**** determinants of an LT-B subunit of the heat-labile enterotoxin of Escherichia coli (LT-BNT).

CLMS(2)

2. The conjugate of claim 1, wherein the nontoxic polypeptide is produced by an Escherichia coli bacterium that has been deposited with the NRRL and assigned accession No. B-15757, or by a mutant, recombinant, or genetically engineered equivalent derivative thereof.

CLMS(3)

3. The conjugate of claim 1, prepared by the ****reductive**** ****amination**** of the PRP fragment and protein.

CLMS(4)

4. The conjugate of claim 1, prepared by ****reductive**** ****amination**** in the presence of cyanoborohydride anions.

CLMS(5)

5. The conjugate of claim 1, wherein said PRP fragment elutes from a column of Bio-Gel P-10 at a V_e/V_o ratio of .1 to .9.

CLMS(6)

6. The conjugate of claim 1, wherein said PRP fragment elutes from a column of Bio-Gel P-10 at a V_e/V_o ratio of 1.09-1.38.

CLMS(7)

7. The conjugate of claim 1 wherein said PRP fragment elutes from a column of Bio-Gel P-10 at a V_e/V_o ratio of 1.39-1.99.

CLMS(8)

8. The conjugate of claim 1, wherein said PRP fragment elutes from a column of Bio-Gel P-10 at a V_e/V_o ratio of 2.0-2.4.

CLMS(9)

9. A vaccine that elicits effective levels of anti-PRP antibody formations in young warm-blooded mammals comprising an ****immunogenic**** amount of the conjugate of claim 5 and a pharmaceutically acceptable carrier.

CLMS(10)

10. A vaccine that elicits effective levels of anti-PRP antibody formations in young warm-blooded mammals comprising an ****immunogenic**** amount of the conjugate of claim 6 and a pharmaceutically acceptable carrier.

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CLAIMS:

CLMS(1)

We claim:

1. A method for actively immunizing human infants against a ****bacterial**** pathogen having a ****capsular**** ****polymer****, comprising:
(a) administering to a human infant an effective amount of an ****immunogenic****-conjugate vaccine comprising a ****capsular**** ****polymer**** or fragment thereof, which is ****immunogenic**** in mature humans but less so in young humans, derived from a ****bacterial**** pathogen selected from the group consisting of Haemophilus influenzae type b, Escherichia coli, Pseudomonas aeruginosa, Neisseria meningitidis and Streptococcus pneumoniae, covalently attached to a ****bacterial**** outer membrane protein or to a ****bacterial**** ****toxin****, ****toxoid**** or binding subunit thereof; and
(b) subsequently administering to said human infant an effective amount of the corresponding intact unconjugated ****capsular**** ****polymer****.

CLMS(2)

2. The method of claim 1, wherein the immunogenic-conjugate vaccine comprises a ****capsular**** ****polymer**** which is ****immunogenic**** in adult humans but not in young humans, derived from the said ****bacterial**** pathogen covalently attached to a ****bacterial**** outer membrane protein or to a ****bacterial**** ****toxin****, ****toxoid**** or binding subunit therefrom.

CLMS(3)

3. The method of claim 1, wherein the ****immunogenic****-conjugate vaccine comprises a ****capsular**** ****polymer**** fragment which is ****immunogenic**** in adult humans but not in young humans, derived from the said ****bacterial**** pathogen covalently attached to a ****bacterial**** outer membrane protein or to a ****bacterial**** ****toxin****, ****toxoid**** or binding subunit therefrom.

CLMS(4)

4. A method for actively immunizing human infants against a **bacterial** pathogen having a **capsular** **polymer** comprising:
(a) administering to a human infant an effective amount of an **immunogenic**-conjugate vaccine, comprising the **reductive** **amination** product of a **capsular** **polymer** or fragment thereof, which is **immunogenic** in mature humans but less so in young humans, having a reducing end and derived from the **capsular** **polymer** of a **bacterial** pathogen selected from the group consisting of Haemophilus influenzae type b, Escherichia coli, Pseudomonas aeruginosa, Neisseria meningitidis and Streptococcus pneumoniae, and a **bacterial** outer membrane protein or a **bacterial** **toxin**, **toxoid** or binding subunit therefrom, and
(b) subsequently administering to said human infant an effective amount of the corresponding intact unconjugated **capsular** **polymer**.

CLMS(5)

5. The method of claim 4, wherein the **immunogenic**-conjugate vaccine comprises the **reductive** **amination** product of an **immunogenic** **capsular** **polymer** having a reducing end and derived from the **capsular** **polymer** of the said **bacterial** pathogen and a **bacterial** outer membrane protein or a **bacterial** **toxin**, **toxoid** or binding subunit therefrom.

CLMS(6)

6. The method of claim 4, wherein the **immunogenic**-conjugate vaccine comprises the **reductive** **amination** product of an **immunogenic** **capsular** **polymer** fragment having, a reducing end and derived from the **capsular** **polymer** of the said **bacterial** pathogen and a **bacterial** outer membrane protein or a **bacterial** **toxin**, **toxoid** or binding subunit therefrom.

CLMS(7)

7. The method of claim 1 or 4, wherein the **capsular** **polymer** is from Haemophilus influenzae type b.

CLMS(8)

8. The method of claim 1 or 4, wherein the **capsular** **polymer** is from Escherichia coli.

CLMS(9)

9. The method of claim 1 or 4, wherein the **capsular** **polymer** is from Neisseria meningitidis serogroup A.

CLMS(10)

10. The method of claim 1 or 4, wherein the unconjugated **capsular** **polymer** is from Neisseria meningitidis serogroup A.

CLMS(11)

11. The method of claim 1 or 4, wherein the **capsular** **polymer** is from Neisseria meningitidis serogroup C.

CLMS(12)

12. The method of claim 1 or 4, wherein the **capsular** **polymer** is from Streptococcus pneumoniae.

CLMS(13)

13. The method of claim 1 or 4, wherein the ****capsular**** ****polymer**** is from *Streptococcus pneumoniae* serotype 3.

CLMS(14)

14. The method of claim 1 or 4, wherein the ****capsular**** ****polymer**** is from *Streptococcus pneumoniae* serotype 6.

CLMS(15)

15. The method of claim 1 or 4, wherein the ****capsular**** ****polymer**** is from *Streptococcus pneumoniae* serotype 12.

CLMS(16)

16. The method of claim 1 or 4, wherein the ****capsular**** ****polymer**** is from *Streptococcus pneumoniae* serotype 14.

CLMS(17)

17. The method of claim 1 or 4, wherein the ****capsular**** ****polymer**** is from *Streptococcus pneumoniae* serotype 19.

CLMS(18)

18. The method of claim 1 or 4, wherein the ****capsular**** ****polymer**** is from *Streptococcus pneumoniae* serotype 23.

CLMS(19)

19. The method of claim 1 or 4, wherein the ****capsular**** ****polymer**** is from *Streptococcus pneumoniae* serotype 51.

CLMS(20)

20. The method of claim 1 or 4, wherein the ****capsular**** ****polymer**** is from *Pseudomonas aeruginosa*.

CLMS(21)

21. The method of claim 1 or 4, wherein the ****toxin****, ****toxoid**** or binding subunit therefrom is from a diphtheria bacterium.

CLMS(22)

22. The method of claim 1 or 4, wherein the ****toxoid**** is diphtheria CRM.

CLMS(23)

23. The method of claim 1 or 4, wherein the ****toxoid**** is diphtheria CRM.sub.197.

CLMS(24)

24. The method of claim 1 or 4, wherein the ****toxin****, ****toxoid**** or binding subunit therefrom is from a tetanus bacterium.

CLMS(25)

25. The method of claim 1 or 4, wherein the ****toxin**** or ****toxoid**** is from a *pseudomonas* bacterium.

CLMS(26)

26. The method of claim 1 or 4, wherein the ****toxin**** or ****toxoid**** is

from a staphylococcus bacterium.

CLMS(27)

27. The method of claim 1 or 4, wherein the ****toxin**** or ****toxoid**** is from a streptococcus bacterium.

CLMS(28)

28. The method of claim 1 or 4, wherein the ****toxin**** or ****toxoid**** is from a pertussis bacterium.

CLMS(29)

29. The method of claim 1 or 4 wherein the outer membrane protein is from Haemophilus influenzae type b.

CLMS(30)

30. The method of claim 1 or 4 wherein the outer membrane protein is from Neisseria meningitidis.

CLMS(31)

31. The method of claim 1 or 4 wherein the outer membrane protein is from Streptococcus pneumoniae.

CLMS(32)

32. The method of claim 1 or 4 wherein the outer membrane protein is from E. coli.

CLMS(33)

33. The method of claim 1 or 4 wherein the outer membrane protein is from a pertussis bacterium.

US PAT NO: 4,761,283

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CLAIMS:

CLMS(1)

I claim:

1. An ****immunogenic**** conjugate, comprising: the ****reductive**** ****amination**** product of a ****capsular**** ****polymer**** fragment having a reducing end and derived from the ****capsular**** ****polymer**** of a ****bacterial**** pathogen selected from the group consisting of Haemophilus influenzae type b, Escherichia coli, Neisseria meningitidis and Streptococcus pneumoniae, and the diphtheria ****toxin**** protein CRM.sub.197.

CLMS(2)

2. The ****immunogenic**** conjugate of claim 1, wherein the ****capsular**** ****polymer**** is ****immunogenic**** in mature humans and less ****immunogenic**** in infant humans.

CLMS(3)

3. The ****immunogenic**** conjugate of claim 1, wherein the ****reductive**** ****amination**** is performed in the presence of cyanoborohydride anions.

CLMS(4)

4. The **immunogenic** conjugate of claim 1, wherein the **bacterial** pathogen is Haemophilus influenzae type b.

CLMS(5)

5. The **immunogenic** conjugate of claim 1, wherein the **bacterial** pathogen is Escherichia coli.

CLMS(6)

6. The **immunogenic** conjugate of claim 1, wherein the **bacterial** pathogen is Neisseria meningitidis.

CLMS(7)

7. The **immunogenic** conjugate of claim 1, wherein the **bacterial** pathogen is Neisseria meningitidis serogroup A.

CLMS(8)

8. The **immunogenic** conjugate of claim 1, wherein the **bacterial** pathogen is Neisseria meningitidis serogroup C.

CLMS(9)

9. The **immunogenic** conjugate of claim 1, wherein the **bacterial** pathogen is Streptococcus pneumoniae.

CLMS(10)

10. The **immunogenic** conjugate of claim 1, wherein the **bacterial** pathogen is Streptococcus pneumoniae serotype 3.

CLMS(11)

11. The **immunogenic** conjugate of claim 1, wherein the **bacterial** pathogen is Streptococcus pneumoniae serotype 6.

CLMS(12)

12. The **immunogenic** conjugate of claim 1, wherein the **bacterial** pathogen is Streptococcus pneumoniae serotype 12.

CLMS(13)

13. The **immunogenic** conjugate of claim 1, wherein the **bacterial** pathogen is Streptococcus pneumoniae serotype 14.

CLMS(14)

14. The **immunogenic** conjugate of claim 1, wherein the **bacterial** pathogen is Streptococcus pneumoniae serotype 19.

CLMS(15)

15. The **immunogenic** conjugate of claim 1, wherein the **bacterial** pathogen is Streptococcus pneumoniae serotype 23.

CLMS(16)

16. The **immunogenic** conjugate of claim 1, wherein the **bacterial** pathogen is Streptococcus pneumoniae serotype 51.

CLMS(17)

17. The **immunogenic** conjugate of claim 1, wherein the fragment is

derived from the **capsular** **polymer** by oxidative cleavage.

CLMS(18)

18. The **immunogenic** conjugate of claim 1, wherein the fragment is derived from the **capsular** **polymer** by periodate.

CLMS(19)

19. The **immunogenic** conjugate of claim 1, wherein the fragment is derived from the **capsular** **polymer** by hydrolysis of a glycosidic linkage.

CLMS(20)

20. The **immunogenic** conjugate of claim 19, wherein the hydrolysis is accomplished enzymatically.

CLMS(21)

21. The **immunogenic** conjugate of claim 19, wherein the hydrolysis is accomplished chemically.

CLMS(22)

22. The **immunogenic** conjugate of claim 19, wherein the hydrolysis is accomplished by acid.

CLMS(23)

23. The **immunogenic** conjugate of claim 4, wherein the fragment elutes on a column of Bio-Gel P-10 at a V_e/V_o ratio of <1.08 .

CLMS(24)

24. The **immunogenic** conjugate of claim 4, wherein the fragment elutes on a column of Bio-Gel P-10 at a V_e/V_o ratio of 1.09-1.38.

CLMS(25)

25. The **immunogenic** conjugate of claim 4, wherein the fragment elutes on a column of Bio-Gel P-10 at a V_e/V_o ratio of 1.39-1.99.

CLMS(26)

26. The **immunogenic** conjugate of claim 14 wherein the fragment elutes on a column of Bio-Gel P-10 at a V_e/V_o ratio of 2.0-2.4.

CLMS(27)

27. An **immunogenic** conjugate, comprising: a formalin treated **reductive** **amination** product of a **capsular** **polymer** fragment having a reducing end and derived from the **capsular** **polymer** of a **bacterial** pathogen selected from the group consisting of Haemophilus influenzae type b, Escherichia coli, Neisseria meningitidis and Streptococcus pneumoniae, and the diphtheria **toxin** protein CRM.sub.197.

CLMS(28)

28. A vaccine that elicits effective levels of anti-**capsular** **polymer** antibodies in humans, comprising: the **immunogenic** conjugate of claim 1.

CLMS(29)

29. A method for actively immunizing humans ****bacterial**** pathogen having a ****capsular**** ****polymer****, comprising: administering an effective amount of the vaccine of claim 28.

CLMS(30)

30. An ****immunogenic**** conjugate of (1) a ****bacterial**** ****capsular**** ****polymer**** fragment having a reducing end, said fragment produced by selective acid hydrolysis of a ****capsular**** ****polymer**** obtained from a ****bacterial**** pathogen selected from the group consisting of selected from the group consisting of Haemophilus influenzae type b, Escherichia coli, Neisseria meningitidis and Streptococcus pneumoniae, without significant destruction of ****antigenic**** specificity, and (2) the diphtheria ****toxin**** protein CRM.sub.197.

CLMS(31)

31. The ****immunogenic**** conjugate of claim 30, wherein the ****capsular**** ****polymer**** is derived from Streptococcus pneumoniae serotype 6 or 12.

CLMS(32)

32. A vaccine that elicits effective levels of anti-polyribosyl ribitol phosphate antibody formations in young warm-blooded mammals comprising an ****immunogenic**** amount of the conjugate of claim 1 and a pharmaceutically acceptable carrier.

CLMS(33)

33. A vaccine that elicits effective levels of anti-polyribosyl ribitol phosphate antibody formations in young warm-blooded mammals comprising an ****immunogenic**** amount of the conjugate of claim 4 and a pharmaceutically acceptable carrier.

CLMS(34)

34. A method for inducing active immunization against systemic infection in young warm-blooded mammals caused by the pathogen Haemophilus influenzae type b comprising administering an ****immunogenic**** amount of the conjugate of claim 4.

US PAT NO: 4,673,574

L6: 6 of 6

CLAIMS:

CLMS(1)

I claim:

1. ****immunogenic**** conjugate comprising the reductive amination product of an ****immunogenic**** ****capsular**** ****polymer**** fragment having a chain length of from about 10 to about 30 monomeric units and a reducing end, which fragment is derived from the ****capsular**** ****polymer**** of a Streptococcus pneumoniae or Haemophilus influenzae bacterium, and a ****bacterial**** ****toxin**** or ****toxoid****.

CLMS(2)

2. The ****immunogenic**** conjugate of claim 1, wherein the ****capsular**** ****polymer**** is ****immunogenic**** in mature humans and less ****immunogenic**** in infant humans.

CLMS(3)

3. The ****immunogenic**** conjugate of claim 1, wherein the ****reductive**** ****amination**** is performed in the persence of cyanoborohydride anions.

CLMS(4)

4. The **immunogenic** conjugate of claim 1, wherein the **toxin** or **toxoid** is diphtheria **toxin** or **toxoid**.

CLMS(5)

5. The **immunogenic** conjugate of claim 4, wherein the **toxoid** is CRM.sub.197.

CLMS(6)

6. The **immunogenic** conjugate of claim 1, wherein the **toxin** or **toxoid** is tetanus **toxin** or **toxoid**.

CLMS(7)

7. The **immunogenic** conjugate of claim 1, wherein the **toxin** or **toxoid** is a pseudomonas **toxin** or **toxoid**.

CLMS(8)

8. The **immunogenic** conjugate of claim 1, wherein the **toxin** or **toxoid** is a staphylococcus **toxin** or **toxoid**.

CLMS(9)

9. The **immunogenic** conjugate of claim 1, wherein the **toxin** or **toxoid** is a streptococcus **toxin** or **toxoid**.

CLMS(10)

10. The **immunogenic** conjugate of claim 1, wherein the **toxin** or **toxoid** is pertussis **toxin** or **toxoid**.

CLMS(11)

11. The **immunogenic** conjugate of claim 1, wherein the **toxin** or **toxoid** is Escherichia coli **toxin** or **toxoid**.

CLMS(12)

12. The **immunogenic** conjugate of claim 1, wherein the **bacterial** pathogen is Haemophilus influenzae type b.

CLMS(13)

13. The **immunogenic** conjugate of claim 1, wherein the **bacterial** pathogen is Streptococcus pneumoniae serotype 3.

CLMS(14)

14. The **immunogenic** conjugate of claim 1, wherein the **bacterial** pathogen is Streptococcus pneumoniae serotype 6.

CLMS(15)

15. The **immunogenic** conjugate of claim 1, wherein the **bacterial** pathogen is Streptococcus pneumoniae serotype 12.

CLMS(16)

16. The **immunogenic** conjugate of claim 1, wherein the **bacterial** pathogen is Streptococcus pneumoniae serotype 14.

CLMS(17)

17. The **immunogenic** conjugate of claim 1, wherein the **bacterial** pathogen is *Streptococcus pneumoniae* serotype 19.

CLMS(18)

18. The **immunogenic** conjugate of claim 1, wherein the **bacterial** pathogen is *Streptococcus pneumoniae* serotype 23.

CLMS(19)

19. The **immunogenic** conjugate of claim 1, wherein the **bacterial** pathogen is *Streptococcus pneumoniae* serotype 51.

CLMS(20)

20. The **immunogenic** conjugate of claim 5, wherein the **bacterial** pathogen is *Haemophilis influenzae* type b.

CLMS(21)

21. The **immunogenic** conjugate of claim 5, wherein the **bacterial** pathogen is *Streptococcus pneumoniae* serotype 6.

CLMS(22)

22. The **immunogenic** conjugate of claim 5, wherein the **bacterial** pathogen is *Streptococcus pneumoniae* serotype 14.

CLMS(23)

23. The **immunogenic** conjugate of claim 5, wherein the **bacterial** pathogen is *Streptococcus pneumoniae* serotype 19.

CLMS(24)

24. The **immunogenic** conjugate of claim 5, wherein the **bacterial** pathogen is *Streptococcus pneumoniae* serotype 23.

CLMS(25)

25. The **immunogenic** conjugate of claim 1, wherein the fragment is derived from the **capsular** **polymer** by oxidative cleavage.

CLMS(26)

26. The **immunogenic** conjugate of claim 1, wherein the fragment is derived from the **capsular** **polymer** by periodate.

CLMS(27)

27. The **immunogenic** conjugate of claim 1, wherein the fragment is derived from the **capsular** **polymer** by hydrolysis of a glycosidic linkage.

CLMS(28)

28. The **immunogenic** conjugate of claim 27, wherein the hydrolysis is accomplished enzymatically.

CLMS(29)

29. The **immunogenic** conjugate of claim 27, wherein the hydrolysis is accomplished chemically.

CLMS(30)

30. The ****immunogenic**** conjugate of claim 27, wherein the hydrolysis is accomplished by acid.

CLMS(31)

31. The ****immunogenic**** conjugate of claim 12, wherein the fragment elutes on a column of Bio-Gel P-10 at a V_e/V_o ratio of .1 to req.1.08.

CLMS(32)

32. The ****immunogenic**** conjugate of claim 12, wherein the fragment elutes on a column of Bio-Gel P-10 at a V_e/V_o ratio of 1.09-1.38.

CLMS(33)

33. The ****immunogenic**** conjugate of claim 12, wherein the fragment elutes on a column of Bio-Gel P-10 at a V_e/V_o ratio of 1.39-1.99.

CLMS(34)

34. An ****immunogenic**** conjugate comprising a formalin treated ****reductive**** ****amination**** product of an ****immunogenic**** ****capsular**** ****polymer**** fragment having a chain length of from about 10 to about 30 monomeric units and a reducing end, which fragment is derived from the ****capsular**** ****polymer**** of a Streptococcus pneumoniae or Haemophilus influenzae bacterium, and a ****bacterial**** ****toxin**** or ****toxoid****.

CLMS(35)

35. The ****immunogenic**** conjugate of claim 34, wherein the ****bacterial**** ****toxoid**** is diphtheria ****toxoid****.

CLMS(36)

36. The ****immunogenic**** conjugate of claim 35, wherein the ****Toxoid**** is CRM.sub.197.

CLMS(37)

37. The ****immunogenic**** conjugate of claim 34, wherein the ****bacterial**** ****toxin**** or ****toxoid**** is tetanus ****toxin**** or ****toxoid****.

CLMS(38)

38. An ****immunogenic**** conjugate of (1) a PRP polysaccharide fragment having reducing terminal groups derived from the capsular polysaccharide of Haemophilus influenzae type b by selective acidic hydrolysis of a portion of the ribosyl ribitol linkages therein and (2) the diphtheria ****toxin**** protein CRM.sub.197.

CLMS(39)

39. The conjugate of claim 38 prepared by the ****reductive**** ****amination**** of the PRP fragment and protein.

CLMS(40)

40. The conjugate of claim 38 prepared by ****reductive**** ****amination**** in the presence of cyanoborohydride anions.

CLMS(41)

41. The conjugate of claim 38 wherein said PRP fragment elutes from a column of Bio-Gel P-10 at a V_e/V_o ratio of .1 to req.1.08.

CLMS(42)

42. The conjugate of claim 38 wherein said PRP fragment elutes from a column of Bio-Gel P-10 at a V_e/V_o ratio of 1.09-1.38.

CLMS(43)

43. The conjugate of claim 38 wherein said PRP fragment elutes from a column of Bio-Gel P-10 at a V_e/V_o ratio of 1.39-1.99.

CLMS(44)

44. The conjugate of claim 38 wherein said PRP fragment elutes from a column of Bio-Gel P-10 at a V_e/V_o ratio of 2.0-2.4.

CLMS(45)

45. A vaccine that elicits effective levels of anti-**capsular** **polymer** antibodies in humans, comprising: the **immunogenic** conjugate of claim 1.

CLMS(46)

46. A method for actively immunizing humans against a **bacterial** pathogen having a **capsular** **polymer**, comprising: administering an effective amount of the vaccine of claim 45.

CLMS(47)

47. A vaccine that elicits effective levels of anti-PRP antibody formations in young warm-blooded mammals comprising an **immunogenic** amount of the conjugate of claim 41 and a pharmaceutically acceptable carrier.

CLMS(48)

48. A vaccine that elicits effective levels anti-PRP antibody formations in young warm-blooded mammals comprising an **immunogenic** amount of the conjugate of claim 42 and a pharmaceutically acceptable carrier.

CLMS(49)

49. The **immunogenic** conjugate of claim 5, wherein the **bacterial** pathogen is *Streptococcus pneumoniae* serotype 3.

CLMS(50)

50. The **immunogenic** conjugate of claim 5, wherein the **bacterial** pathogen is *Streptococcus pneumoniae* serotype 51.

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d his

(FILE 'CA' ENTERED AT 14:30:26 ON 30 SEP 92)
DEL HIS

L1	11	S	CAPSULAR POLYMER#
L2	1767	S	REDUCTIVE AMINATION#
L3	1281	S	BACTERIAL (10A) (TOXIN# OR TOXOID#)
L4	0	S	L1 AND L2 AND L3
L5	140618	S	LINK? OR CROSSLINK? OR CONJUGAT?
L6	3	S	L1 AND L3 AND L5
L7	23214	S	POLYSACCHARIDE#
L8	0	S	L1 AND L2
L9	80	S	REDUCTIVE DEAMINATION#
L10	0	S	L1 AND L2 AND L3
L11	2	S	L7 AND L9
L12	0	S	L1 AND L9 AND L3

=>

File CA 81930
File CA 819305

```

capsular polymer#
      1159 CAPSULA
      447413 POLYMER#
L1      11 CAPSULAR POLYMER#
      (CAPSULAR(W)POLYMER#)

=> s reductive amination#
      19126 REDUCTIVE
      14957 AMINATION#
L2      1767 REDUCTIVE AMINATION#
      (REDUCTIVE(W)AMINATION#)

=> s bacterial (00a) toxin# or toxoid#
      50397 BACTERIAL
      32371 TOXIN#
      1178 TOXOID#
L3      1281 BACTERIAL (10A) (TOXIN# OR TOXOID#)

=> s 11 and 12 and 13
L4      0 L1 AND L2 AND L3

=> s link? or crosslink? or conjugat?
      26923 LINK?
      81393 CROSSLINK?
      37082 CONJUGAT?
L5      140618 LINK? OR CROSSLINK? OR CONJUGAT?

=> s 11 and 13 and 15
L6      3 L1 AND L3 AND L5

=> d 1-3

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L6  ANSWER 1 OF 3  COPYRIGHT 1992 ACS
AN  CA110(14):121374y
TI  Bacterial polymer capsule-bacterial toxin conjugate vaccines for
    infant immunization
AU  Anderson, Porter W.
CS  University of Rochester
LO  USA
SO  U.S., 14 pp. Cont.-in-part of U.S. 4,673,574.
PI  US 4762713 A 9 Aug 1988
AI  US 85-732200 8 May 1985
PRAI US 81-298102 31 Aug 1981
    US 83-511048 5 Jul 1983
IC  H61K039-02; H61K039-095
NCL 424092000
SC  63-3 (Pharmaceuticals)
SX  15
DT  P
CO  USXXAM
PY  1988
LA  Eng

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L6  ANSWER 2 OF 3  COPYRIGHT 1992 ACS
AN  CA107(18):161671n
TI  Immunogenic conjugates for vaccines against childhood diseases.
AU  Anderson, Porter W.
LO  USA
SO  U.S., 11 pp. Cont.-in-part of U.S. Ser. No. 298,102, abandoned.
PI  US 4673574 A 16 Jun 1987
AI  US 83-511048 5 Jul 1983
PRAI US 81-298102 31 Aug 1981
IC  ICM A61K039-02
    ICS A61K039-09; A61K039-102; C07K015-34
NCL 424092000

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53-3 (Pharmaceuticals)
SX 15
DT P
CO USXXAM
PY 1987
LA Eng

L6 ANSWER 3 OF 3 COPYRIGHT 1992 ACS
AN CA105(18):158789k
TI Immunogenic conjugates of E. coli LT-B enterotoxin subunit and capsular polymers
AU Anderson, Porter W.; Clements, John D.
CS Praxis Biologics, Inc.
LO USA
SO Eur. Pat. Appl., 122 pp.
PI EP 172107 A1 19 Feb 1986
DS R: AT, BE, CH, DE, FR, GB, IT, LI, LU, NL, SE
AI EP 85-401604 6 Aug 1985
PRAI US 84-639293 10 Aug 1984
IC ICM A61K039-02
ICS A61K039-102; A61K039-108; A61K039-095; A61K039-09; A61K039-104; C07K015-00; C12P021-00; C12N015-00; A61K039-385; A61K039-395
ICI C12R001-19, C12R001-38, C12R001-21, C12R001-36, C12R001-46, C12R001-63
SC 63-3 (Pharmaceuticals)
SX 1, 15
DT P
CO EPXXDW
PY 1986
LA Eng

=> s polysaccharide#
L7 23214 POLYSACCHARIDE#

=> s 11 and 12
L8 0 L1 AND L2

=> s reductive deamination#
19126 REDUCTIVE
3270 DEAMINATION#
L9 80 REDUCTIVE DEAMINATION#
(REDUCTIVE(W)DEAMINATION#)

=> s 11 and 12 and 13
L10 0 L1 AND L2 AND L3

=> s 17 and 19
L11 2 L7 AND L9

=> d 1-2

L11 ANSWER 1 OF 2 COPYRIGHT 1992 ACS
AN CA93(17):163628v
TI Some new methods for structural elucidation and modification of complex carbohydrates
AU Loenngren, Joergen
CS Dep. Org. Chem., Univ. Stockholm
LO Stockholm S-106 91, Swed.
SO Int. Congr. Pure Appl. Chem., [Proc.], 27th, 205-11
SC 9-0 (Biochemical Methods)
DT J
CO PCPAAI
IS 0359-0561
PY 1980
LA Eng

AN CA93(15):150539s
 TI Reductive deamination of aminodeoxy groups in glycosides and polysaccharides
 AU Arnarp, Jan; Garegg, Per J.; Lengstad, Bengt; Loenngren, Joergen
 CS Dep. Org. Chem., Univ. Stockholm
 LO Stockholm S-106 91, Swed.
 SO Carbohydr. Res., 83(2), 394-7
 SC 33-5 (Carbohydrates)
 DT J
 CO CRRAT
 IS 0008-6215
 PY 1980
 LA Eng

=> s 11 and 19 and 13
 L12 0 L1 AND L9 AND L3

=>

L11 ANSWER 1 OF 2 COPYRIGHT 1992 ACS

AB A review with 19 refs. The detn. of abs. configuration of monosaccharides using gas chromatog; the detn. of abs. configuration of pyruvic acid acetals present in polysaccharides; the deamination of aminodeoxy sugars; and the structural elucidation of the capsular polysaccharides of *Streptococcus pneumoniae* type 1 were discussed.

L11 ANSWER 2 OF 2 COPYRIGHT 1992 ACS

AB The title deamination was carried out with $\text{H}_2\text{NOS}_3\text{H}$. Deoxyglycosides were obtained in 19-52% yield, e.g., Me 2-amino-2-deoxy-.beta.-D-glucopyranoside hydrochloride gave 52% Me 2-deoxy-.beta.-D-arabino-hexopyranoside. *Streptococcus pneumoniae* Type 14 capsular polysaccharide and *Vibrio cholerae* O-antigen were first N-deacylated and then deaminated to give 55 and 60% resp. partly deaminated polysaccharide.

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